

U.S. DEPARTMENT OF COMMERCE National Technical Information Service

AD-A035 260

PULMONARY FUNCTIONS IN CONSCIOUS AND ANESTHETIZED RHESUS MONKEYS

ARMY MEDICAL RESEARCH INSTITUTE OF INFECTIOUS DISEASES, FREDERICK, MARYLAND

25 JANUARY 1977

UNCLASSIFIED

SECURITY CLASSIFICATION OF THIS PAGE (When Data Entered)

REPORT DOCUMEN		READ INSTRUCTIONS BEFORE COMPLETING FORM
I. REPORT NUMBER	2. GOVT ACCESSION NO.	3. RECIPIENT'S CATALOG NUMBER
4. TITLE (and Substitle) Pulmonary Functions in Consc	cious and Anesthetized	5. Type of REPORT & PERIOD COVERED Interim
Rhesus Monkeys		6. PERFORMING ORG. REPORT NUMBER
7. AUTHOR(*) Ching~Tong Liu and Richard I). DeLauter	8. CONTRACT OR GRANT NUMBER(*)
9. PERFORMING ORGANIZATION NAME AND U.S. Army Medical Research I Infectious Diseases SC Fort Detrick, Frederick, Mar	Institute of GRD-UIA-A	10. PROGRAM ELEMENT, PROJECT, TASK AREA & WORK UNIT NUMBERS 62760A 3M62776A841 00 030
11. CONTROLLING OFFICE NAME AND ADD	PRESS	12. REPORT DATE 25 January 1977 13. NUMBER OF PAGES 26
14. MONITORING AGENCY NAME & ADDRES	S(II different from Controlling Office)	15. SECURITY CLASS. (of this report) UNCLASSIFIED 15a. DECLASSIFICATION/DOWNGRADING SCHEDULE

Approved for public release; distribution unlimited

17. DISTRIBUTION STATEMENT (of the abstract entered in Block 20, if different from Report)

18. SUPPLEMENTARY NOTES

Reprints bearing assigned AD number will be forwarded upon receipt. To be submitted for publication in the American Journal of Veterinary Research.

19. KEY WORDS (Continue on reverse side if necessary and identify by block number)

Macaca mulatta Pulmonary functions Blood gas tension Base-line values Anesthesia

20. ABSTRACT (Continue on reverse side if necessary and identity by block number)

Techniques for measurements of selected pulmonary functions in conscious and anesthetized rhesus macaques (Macaca mulatta) are described, and normal baseline values are presented during periods of respiration using pure 02 or room air. Tidal volume, respiratory rate, minute volume, 0, consumption, and specific ventilation were determined nourly for a period of 5 hours in conscious monkeys. Additional variables including dynamic pulmonary compliance, pulmonary resistance, specific compliance, intraesophageal pressure, transpulmonary pressure, inspiratory air flow, expiratory air flow, physiological (cont'd)

DD 1 JAN 73 1473 EDITION OF 1 NOV 65 IS OBSOLETE

UNCLASSIFIED SECURITY CLASSIFICATION OF THIS PAGE (When Date Entered)

TINICT /	100	TEN	ren
UNCL	122	IF.	LED

SECURITY CLASSIFICATION OF THIS PAGE(When Date Entered)

Item 20 concluded:

dead space, $\rm CO_2$, output, respiratory quotient and functional residual capacity, as well as arterial blood pH, $\rm P_{O_2}$, $\rm P_{CO_2}$, $\rm HCO_3$, and total $\rm CO_2$ were measured in anesthetized macaques. Further, comparisons of values for certain pulmonary functions were made between this and other studies.

10

Pulmonary Functions in Conscious and Anesthetized Rhesus Monkeys

C. T. Liu, PhD, and R. D. DeLauter

Please send proofs to: C. T. Liu, PhD

Research Physiologist Animal Assessment Division USAMRIID, Fort Detrick Frederick, Md 21701 D D C

PROCEDURA

197

LEGETUTEU

A

Approved for public release; distribution unlimited

25 Jan 77

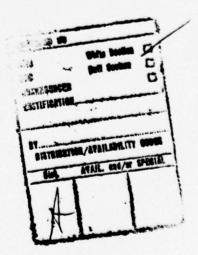
REPRODUCED BY
NATIONAL TECHNICAL
INFORMATION SERVICE
U. S. DEPARTMENT OF COMMERCE
SPRINGFIELD, VA. 22161



Pulmonary Functions in Conscious and Anesthetized Rhesus Monkeys

C. T. Liu, PhD, and R. D. DeLauter

Please send proofs to: C. T. Liu, PhD
Research Physiologist
Animal Assessment Division
USAMRIID, Fort Detrick
Frederick, Md 21701



From the U. S. Army Medical Research Institute of Infectious Diseases, Fort Detrick, Frederick, Maryland 21701

In conducting the research described in this report, the investigators adhered to the "Guide for the Care and Use of Laboratory Animals," as promulgated by the Committee on the Revision of the Guide for Laboratory Animal Facilities and Care of the Institute of Laboratory Animal Resources, National Research Council. The facilities are fully accredited by the American Association for Accreditation of Laboratory Animal Care.

Acknowledgements

The authors thank CPT C. L. Hadick for surgical operations, Mr. D. C. Smith for constructing the monkey head cover, SP5 D. R. Dufault for the drawing, Mrs. Phebe W. Summers for editorial assistance and Mrs. Regina E. Staley for secretarial aid.

SUMMARY

Techniques for measurements of selected pulmonary functions in conscious and anesthetized rhesus macaques (Macaca mulatta) are described, and normal base-line values are presented during periods of respiration using pure 0_2 or room air. Tidal volume, respiratory rate, minute volume, 0_2 consumption, and specific ventilation were determined hourly for a period of 5 hours in conscious monkeys. Additional variables including dynamic pulmonary compliance, pulmonary resistance, specific compliance, intraesophageal pressure, transpulmonary pressure, inspiratory air flow, expiratory air flow, physiological dead space, $C0_2$ output, respiratory quotient and functional residual capacity, as well as arterial blood pH, $P0_2$, $PC0_2$, $HC0_3$ and total $C0_2$ were measured in anesthetized macaques. Further, comparisons of values for certain pulmonary functions were made between this and other studies.

KEY WORD INDEX--Macaca mulatta, pulmonary functions, blood gas tension, base-line values, anesthesia.

INTRODUCTION

Since nonhuman primates are phylogenetically related to man, rhesus and cynomolgus monkeys are commonly used as respiratory models 2,3,8,11,12,15 for studying metabolism, 6,7,23 inhalation toxicity, 13 allergic states, 14,21 asthma, 22 cigarette smoke, 1 and X-irradiation.5 Although base-line values of pulmonary functions in both conscious and anesthetized monkeys have been reported by several investigators, 2-4,8,11, 12,14,15 the data have been incomplete with respect to certain pulmonary functions or the functions were depressed markedly by anesthesia. Further, data variablility as a function of duration of study was not given. The objectives of this study were: (1) to develop techniques for studying selected multiple pulmonary functions in conscious and anesthetized rhesus monkeys, (2) to establish an anesthetized monkey model with minimal depression of pulmonary function, (3) to develop base-line values for major pulmonary functions in anesthetized monkeys and compare them with data obtained by other investigators, (4) to demonstrate the variability of pulmonary data hourly for a period of 5 hours, and (5) to determine if certain respiratory functions are altered by breathing pure $\mathbf{0}_2$ in comparison to room air.

Materials and Methods

Conscious Model Four male rhesus monkeys weighing approximately 3.5 kg were restrained in primate chairs. Before chairing, each monkey was sedated with ketamine HCl (5 mg/kg intramuscularly, im) and the neck hair was closely shaved. After the monkey was completely recovered from effects of ketamine, a head cover (Fig 1) was placed on the top of the chair and the central openings of the soft rubber membranes were fitted properly around the monkey's neck. A thin layer of silicone stopcock grease was applied to the skin of the neck to prevent air leakage from the system. The inlet and outlet processes of the head cover were connected to a 9-liter spirometer (residual volume apparatus) that had been filled with pure 0_2 for continuous breathing. After the monkey was adapted to the chair, head cover, and 02 breathing for 2 hours, tidal volume, respiratory rate, and 0, consumption were determined hourly for a period of 5 hours. At each hour, a 10-15-minute record was obtained. Minute volume (rate x tidal volume) and specific ventilation (minute volume/0, consumption) were calculated, and certain values were also expressed in terms of body surface area. 18

Anesthetized Model A femoral artery and vein were cannulated under halothane [with 0₂ (2%) and maintained at 1.5-1.0%] anesthesia 1 day before measuring pulmonary functions. During an experiment, the monkey was anesthetized again with ketamine (15 mg/kg, im) and a light surgical level of anesthesia was maintained by constant infusion of Na pentobarbital into the femoral vein. The concentration of pentobarbital was (1 mg/cc) and the dose rate was 0.25 mg/minute. The

palpebral reflex was used as an index to determine whether pentobarbital infusion should be continued or discontinued temporarily.

The anesthetized monkey was studied while resting in a supine position. Experiments began 2 hours after completion of endetracheal and intraesophageal intubation. The esophageal tube (i.d. = 6.0 mm, o.d. = 8.5 mm, length = 37.0 cm) was connected to a Statham pressure transducer, and intraesophageal pressure was recorded on a Brush recorder (Fig 2). The intraesophageal pressure was considered as an estimation of intrapleural pressure.

The endotracheal tube was connected sequentially to one of the following devices, while the monkey was breathing room air: (1) pneumotachograph (size 000), (2) 1-way valve or (3) 2-way valve. Tidal volume obtained by electrically integrating the signal from the pneumotachograph and differential pressure transducer, e registering on a Brush recorder. The information on tidal volume and intraesophageal pressure was sent to a digital signal analyzer and stored on magnetic tape. Transpneumotachographic pressure was calculated from the air flow rate during inspiration with known pressure calibration. The difference between measured intraesophageal pressure and calculated transpneumotachographic pressure was defined as transpulmonary pressure.

The 1-way valve was used to separate inspired and expired air, which flowed continuously (50 ml/minute) into a ${\rm CO_2}$ analyzer for ${\rm CO_2}$ concentration (volume %) determination. The 2-way valve was connected to a 9-liter spirometer containing 100% ${\rm O_2}$ for tidal volume, respiratory rate, and ${\rm O_2}$ -consumption determination. Using the same 2-way valve and spirometer, functional residual capacity (FRC) was measured by the helium dilution technique.

Dynamic pulmonary compliance and resistance were calculated by the method described by Giles et al¹⁰ except that the present data were calculated manually. Magnetically recorded tidal volume and intraesophageal pressure were redisplayed on a digital signal averager and the magnitude and time for determinations of isovolumetric points were accurately measured. Equations in Table 1 were used to calculate various pulmonary functions, and certain variables were also expressed in terms of m² of body surface area.¹⁸

Two 1-ml blood samples were taken from the femoral artery, when the monkey was breathing pure 0_2 or room air. Arterial blood pH, P_{02} , total CO_2 , HCO_3^- and base excess were determined in a blood gas analyzer. Blood gases and pH values, as well as certain pulmonary variables were compared between breathing 0_2 and room air.

Statistics. An independent t test was used to compare corresponding pulmonary function measurements in conscious and anesthetized monkeys. When comparisons were made hourly in the same monkey during the 5-hour experiment, a paired t test was used. Data obtained by other investigators 5,8,11,12,14,15,23 were presented as mean, standard deviation, range, or 95% confidence limit (1.96 x standard error of the mean). Comparisons between the present and others' data on pulmonary functions in the anesthetized rhesus monkeys were based upon whether the present data mean values fell within or outside the range of 95% confidence limits of the data of other investigators.

Results

Normal values for various pulmonary functions, including lung volumes, ventilation, and breathing mechanics of anesthetized rhesus monkeys breathing room air, are summarized in Table 2. During a 5-hour period, all measured values varied slightly, except for respiratory rate, which was significantly depressed between 3 and 4 hours. Physiological dead space fluctuated and dynamic pulmonary resistance increased gradually as a function of time; however, these changes were not significant.

Selected pulmonary functions were compared between conscious and anesthetized monkeys during a period of 5 hours (Table 3). Among all measured parameters, there were no significant differences between the 2 groups of monkeys studied.

Values for respiratory quotient (RQ) are presented in Table 4. Similarities of expired ${\rm CO}_2$ concentration, ${\rm CO}_2$ output, arterial blood pH, ${\rm P_{CO}}_2$, total ${\rm CO}_2$ and base excess between breathing pure ${\rm O}_2$ and room air are tabulated in Tables 4 and 5. Blood ${\rm P_{O}}_2$, was much higher in monkeys breathing pure ${\rm O}_2$ than room air (Table 5). Values of RQ, blood pH, and gas tensions were maintained in a relatively constant state throughout a 5-hour period.

Data on various pulmonary functions, 0_2 consumption, and RQ in anesthetized rhesus monkeys from this and other studies are summarized in Tables 6 and 7. In general, results from the present study agree well with the findings of others, except that higher values for 0_2 consumption (compared to Rakieten²³) were demonstrated. The present work also showed higher values for tidal volume, respiratory rate, and minute volume as compared to reported data of Kelly et al. ¹⁴ Further, the measured tidal volumes of this study were higher than values obtained by Crosfill and Widdicombe 8 and Guyton. ¹¹

Discussion

1.

Although pulmonary depression is commonly seen in man and animals under anesthesia, respiratory functions of anesthetized monkeys in this study were not modified significantly compared to data obtained from conscious monkeys. However, the anesthetized monkey showed a trend toward an increased dynamic pulmonary resistance as a function of time. Since few respiratory measurements can be carried out in a conscious monkey, the lightly anesthetized monkey seems to be acceptable for establishing normal values for pulmonary functions, 02 consumption, CO2 output and RQ.

Major advantages for using the lightly anesthetized monkey include:

(1) a great variety of pulmonary functions can be measured; (2) results are not influenced by the monkey's emotions and its associated muscular activities; (3) monkeys recover completely after these measurements of pulmonary function; and (4) monkeys may be used for short-term experiments for evaluation of drugs or toxin. When conscious chaired monkeys are used, the head cover provides minimum disturbances or stimulation to the monkey as compared to the use of a face mask. Further, the closed head cover is suitable for continuous application of 02, aerosol, or positive-pressure breathing for the treatment of pulmonary edema. 17

Normal values for arterial blood gases and pH in conscious rhesus monkeys have been reported by Liu, 16 Munson et al, 20 Forsyth and coworkers, 9 and Binns et al. 3 In the present study, arterial blood values for anesthetized monkeys showed slight decreases in pH, P_{02} as compared to several other investigators. 9,20 However, the present data on arterial blood pH, P_{02} and P_{CO_2} agree well with Binns et al 3 and our previous findings in conscious rhesus monkeys. 16 This evidence supports the

observations that monkeys used in this study were not deeply anesthetized.

There were few differences in arterial blood pH, P_{CO_2} , HCO_3^- , total CO_2 , and base excess between breathing pure O_2 or room air in anesthetized monkeys. However, blood P_aO_2 increased from 83-94 mm Hg to 367-411 mm Hg during pure O_2 breathing. These appeared to be the highest values for P_{O_2} in the arterial blood of lightly anesthetized monkeys.

Some unique aspects of the present study may be summarized as follows:

(1) no significant differences of measured respiratory values were found between conscious and anesthetized models; (2) many respiratory and metabolic variables can be measured in a single lightly anesthetized monkey; (3) the measured respiratory values, except dynamic pulmonary resistance and physiological dead space, remained relatively constant throughout a period of 5 hours; and (4) transpulmonary pressure was calculated by the difference of intraesophageal and transpneumotachographic pressures during inspiration.

In general, values for respiratory and metabolic functions agree well with other workers. 5,8,11,12,14,15,23 However, values for $^{0}2$ consumption and tidal volume were higher in this study than those obtained by other investigators. 8,11,14,23 These discrepancies may result from different levels of anesthesia applied to rhesus monkeys.

Footnotes

Warren E. Collins, Inc., Braintree, Ma. (All original hoses were replaced by small Tygon tubing (i.d. = 10 mm, o.d. = 14.6 mm).

b"Argyle" Sherwood Medical Industries, St. Louis, Mo.

^cStatham, P23BB, Hato Rey, Puerto Rico.

^dDynasciences Medical Products, Blue Bell, Pa. 19422.

^eStatham, PM15E, Hato Rey, Puerto Rico.

f Northern NS-575, Northern Scientific, Inc., Middleton, Wi.

gNorthern NS-408F, Northern Scientific, Inc., Middleton, Wi.

hModel 2050, Harvard Apparatus Co., Millis, Ma.

ⁱCorning model 165, Corning Scientific Instruments, Medfield, Ma.

TABLE 1--Equations for Calculating Pulmonary Functions

	Variable	Equation
ו פי	Physiological dead space $(v_{\overline{D}})^*$	$V_{D} (m1) = (P_{a}CO_{2} - P_{E}CO_{2}) \cdot V_{T}/P_{a}CO_{2}$
79	Functional residual capacity (FRC)**	FRC (m1) = $V_1 [(N_1 - N_2)/N_2]$
Q	CO ₂ output	CO_2 output (ml/min) = minute vol (ml/min) · CO_2 conc. (vol %) in the expired air.
Ð	Dynamic pulmonary compliance (DPC)	DPC $(m1/cm H_20) = \frac{Tidal \ volume \ (m1)}{Transpulmonary \ pressure \ (cm H_20)}$
Ð	Dynamic pulmonary resistance (DPR)	DPR (cm $H_2O/L/sec$) = $(P_1 - P_2)/(F_1 + F_2)$.
×	Respiratory quotient (RQ)	$RQ = CO_2$ output $(m1/min)/O_2$ consumption $(m1/min)$
* *	*P _a CO ₂ = PCO ₂ in arterial blood (mm Hg), P_E CO ₂ = PCO ₂ in expired air (mm Hg), V_T = Tidal vol (ml). **N ₁ (vol %) = Helium (He) concentration before connecting to the monkey, V_1 = vol of He (250 ml)	V_2 in expired air (mm Hg), V_T = Tidal vol (ml). The secting to the monkey, V_{τ} = vol of He (250 ml)
*	** N_1 (vol %) = Helium (He) concentration before connecting to the monkey, V_1 = vol of He (250 ml)	necting to the monkey, V_1 = vol of He (250 ml)

and air in the spirometer = (250 ml/N $_1$), N $_2$ = He concentration after connecting to the animal at the end of expiration.

 P_1 and P_2 are intraesophageal pressure levels at isovolumetric points of tidal volume, F_1 and F_2 are inspiratory and expiratory flow levels at isovolumetric points of tidal volume.

TABLE 2--Pulmonary Functions in Anesthetized Normal Rhesus Monkeys Breathing Room Air

Classification	Variable			Value at time in hours	in hours	
		1	2	3	7	5
Lung volume	Tidal volume (ml)	35 ± 4	37 ± 4	36 ± 4	41 ± 7	41 ± 6
	Tidal volume (ml/m^2L)	129 ± 11	139 ± 15	135 ± 16	152 ± 20	153 ± 21
	FRC (m1)	119 ± 12	108 + 8	111 ± 7	113 ± 10	112 ± 9
	FRC (m1/m ²)	77 + 757	408 + 26	419 + 24	423 + 30	426 ± 36
Pulmonary ventilation	Respiratory rate (cycle/min)	39 + 2	35 + 1	28 + 2*	28 + 2*	35 + 2
	Minute volume (liter/min)	1.36 ± 0.16	1.30 ± 0.16	0.98 ± 0.13	1.21 ± 0.23	1.44 ± 0.26
	Minute volume $(m1/min/m^2)$	4.99 + 0.4	4.84 ± 0.5	3.74 ± 0.5	4.43 ± 0.8	5.26 ± 0.7
	Physiological dead space (ml)	4.33 ± 1.97	0.64 + 3.65	2.52 ± 2.44	0.97 ± 3.9	0.45 ± 3.9
Mechanics of breathing	Intraesophageal pressure (cm $_{ m H_2^{0}}$)	-5.7 ± 0.6	-6.6 ± 0.7	-5.6 ± 0.8	-6.5 + 0.8	-7.1 ± 0.9
	Transpneumotach pressure (cm $_{ m H_2^0}$)	1.13 ± 0.09	1.05 ± 0.10	0.85 ± 0.07	0.99 ± 0.12	1.05 ± 0.12
	Transpulmonary pressure (cm H_2 0)	-4.54 ± 0.55	-5.58 ± 0.67	-4.75 ± 0.78	-5.50 ± 0.76	-6.03 ± 0.91
	Expiratory flow $(m1/cm H_2^0)$	66.6 ± 6.1	0.6 + 0.59	65.0 ± 9.0 56.9 ± 8.7	58.6 ± 7.8	61.9 ± 7.5

TABLE 2--Continued

56.1 ± 6.6 59.5 ± 6.8	8.6 ± 1.7 7.9 ± 1.7	40.9 ± 10	52 ± 7
	8.6 ± 1.7	33.2 ± 5.7	75 ± 13
64.0 ± 5.1 59.5 ± 5.9 48.4 ± 3.7	8.2 ± 0.9 7.3 ± 1.1 9.4 ± 1.8	19.1 ± 5.2 31.3 ± 7.9 32.7 ± 7.8 33.2 ± 5.7 40.9 ± 10	83 ± 14
59.5 ± 5.9	7.3 ± 1.1	31.3 ± 7.9	6 + 99
64.0 ± 5.1	8.2 ± 0.9	19.1 ± 5.2	31 72 ± 9
<pre>Inspiratory flow (m1/sec)</pre>	Dynamic pulmonary compliance (m1/cm H ₂ 0)	Dynamic pulmonary resistance (cm $H_2^0/liter/sec$)	Specific compliance [(compliance/FRC)10-3]

All values are means \pm 1 standard error of the mean (n=9). *P < 0.05.

1

TABLE 3--Comparisons of Selected Pulmonary Functions between Conscious and Anesthetized* Rhesus Monkeys while Breathing $\boldsymbol{\theta}_2$

Variable	Group (n)		Δ	Value by time in hours	hours	
			3	7	5	9
Tidal volume	Conscious	9 + 07	41 + 5	9 + 07	9 + 07	9 + 07
(m1)	Anesthetized	33 + 3	. 38 ± 4	41 + 4	42 + 3	42 + 3
Tidal volume	Conscious	164 ± 30	169 ± 27	164 + 30	164 ± 30	164 ± 30
(m1/m ²)	Anesthetized	124 ± 10	142 ± 11	155 ± 14	157 ± 12	157 ± 11
Respiratory rate	Conscious	28 ± 3	27 ± 3	27 ± 3	25 ± 2	24 + 3
(cycle/min)	Anesthetized	27 ± 1	24 ± 1	21 ± 2	22 ± 1	21 + 2
Minute volume	Conscious	1.10 ± 0.15	1.08 ± 0.14	1.30 ± 0.14	0.99 ± 0.14	0.94 ± 0.12
(liter/min)	Anesthetized	0.90 + 0.08	0.89 ± 0.05	0.81 ± 0.03	0.86 ± 0.05	0.85 + 0.05
Minute volume	Conscious	4.62 ± 0.73	4.54 ± 0.61	4.34 ± 0.62	4.16 ± 0.64	3.97 ± 0.53
(liter/min/m ²)	Anesthetized	3.34 ± 0.25	3.37 ± 0.18	3.11 ± 0.20	3.43 ± 0.15	3.24 ± 0.25
0 ₂ consumption	Conscious	3.56 ± 0.32	3.49 ± 0.49	3.33 ± 0.41	2.84 ± 0.27	3.02 ± 0.53
(liter/hr)	Anesthetized	3.55 ± 0.48	3.95 ± 0.45	3.12 ± 0.38	3.65 ± 0.30	3.14 ± 0.24
0 ₂ consumption	Conscious	14.7 ± 1.0	14.5 ± 1.7	13.8 ± 1.3	11.8 ± 0.8	12.4 ± 1.7
$(1iter/hr/m^2)$	Anesthetized	13.5 ± 1.8	15.0 ± 1.6	12.1 ± 1.5	13.8 ± 0.7	12.1 ± 1.7
Specific ventila-	Conscious	17.8 ± 1.7	18.0 ± 1.1	20.1 ± 2.2	18.1 ± 2.0	19.0 ± 2.8
tion (minute vol/ Anesthetized	Anesthetized	16.3 ± 1.9	14.2 ± 1.1	16.3 ± 1.2	15.1 ± 0.8	16.6 ± 1.4
0 ₂ consumption)						

*Anesthesia: 15 mg/kg ketamine (IM) + 0.25 mg/min of Na pentobarbital (intermittent IV infusion). All values are means + 1 standard error of the mean: n=4 for conscious, n=9 for anesthetized.

TABLE 4--RQ Values and Comparisons of Expired ${\rm CO_2}$ Concentration and ${\rm CO_2}$ Output between Breathing Pure ${\rm O_2}$ and Room Air in Anesthetized Rhesus Monkeys

						-
Variable	Gas	1	2	Value by time in hours	hours 4	5
RQ	Pure 0 ₂	0.80 + 0.08	0.70 ± 0.07	0.82 ± 0.05	0.75 ± 0.05	0.82 ± 0.08
Expired CO ₂ (vol %)	Pure 0_2 Room air	5.03 ± 0.29 4.22 ± 0.13	4.93 ± 0.27 4.26 ± 0.16	5.12 ± 0.23 4.38 ± 0.18	5.00 ± 0.22 4.44 ± 0.13	4.97 ± 0.25 4.21 ± 0.19
<pre>CO₂ output (liter/hr)</pre>	Pure 0_2 Room air	2.45 ± 0.19 3.49 ± 0.40	2.66 ± 0.22 3.39 ± 0.43	$2.50 \pm 0.18 \\ 2.62 \pm 0.36$	2.72 ± 0.18 3.27 ± 0.65	2.54 ± 0.23 3.64 ± 0.71
CO ₂ output (liter/hr/	Pure 0_2 Room air	10.0 ± 0.8 13.0 ± 1.2	10.0 ± 0.7 12.6 ± 1.4	9.6 ± 0.8	10.3 ± 0.6 12.1 ± 2.2	9.7 ± 0.9

All values are means + 1 standard error of the mean.

1

TABLE 5--Comparisons of Arterial Blood pH, Gas Tension HCO_3^- , Total CO_2 , and Base Excess between Breathing Pure O_2 and Room Air in Anesthetized Rhesus Monkeys

Variable	Gas		Value b	Value by time in hours	7	5
Hq	0 ₂ Room air	7.374 ± 0.009	7.364 ± 0.009	7.374 ± 0.009 7.364 ± 0.009 7.362 ± 0.008 7.377 ± 0.005 7.378 ± 0.007 7.374 ± 0.010	7.369 ± 0.009 7.372 ± 0.011	7.369 ± 0.009 7.365 ± 0.010 7.372 ± 0.011 7.376 ± 0.011
PO ₂ (mm Hg)	0 ₂ Room air	367 ± 32 83.2 ± 4.2	370 ± 27 88.7 ± 7.5	389 <u>+</u> 23 88.8 <u>+</u> 3.6	407 ± 12 93.7 ± 5.4	411 ± 16 88.9 ± 3.6
PCO ₂ (mm Hg)	02 Room air	33.4 ± 2.4 30.2 ± 2.5	31.6 ± 3.0 28.2 ± 3.1	33.0 ± 2.8 30.2 ± 2.9	31.5 ± 2.2 28.5 ± 2.6	31.3 ± 2.6 28.0 ± 2.9
HCO ₃ (mM/liter)	0_2 Room air	19.1 ± 1.5 18.5 ± 1.5	18.2 ± 1.9 16.7 ± 1.8	19.0 ± 1.7 18.1 ± 1.7	17.8 ± 1.6 17.1 ± 1.8	18.1 ± 1.7 16.8 ± 1.8
Total CO ₂ (mM/liter)	0 ₂ Room afr	20.4 ± 1.5 20.2 ± 1.6	19.7 ± 1.9 18.1 ± 1.8	20.6 ± 1.7 19.5 ± 1.8	19.5 ± 1.5 18.4 ± 1.8	19.6 <u>+</u> 1.6 18.3 <u>+</u> 1.8
Base excess (mM/liter)	0_2 Room air	-3.7 ± 1.3 -4.2 ± 1.3	-5.1 ± 1.7 -5.7 ± 1.6	-4.2 ± 1.4 -4.8 ± 1.5	-5.0 ± 1.4 -5.6 ± 1.6	-5.0 ± 1.4 -5.7 ± 1.6

All values are means + 1 standard error of the mean (n=9).

TABLE 6--Comparison of Values Obtained by Authors and Other Investigators for Respiratory Rate, Tidal Volume, and Minute Volume in Anesthetized Rhesus Monkeys.

81

No. of animals (wt in kg)	Rate (cycle/min)	Tidal volume (ml)	Minute volume (ml/min)	Reference
9* (3.5 - 6.6)	33.1 ± 5.4 (21 - 48)	38 <u>+</u> 15.1 (12.7 - 79)	1256 <u>+</u> 561 (570 – 2848)	Present study
11 (3.1 - 5.4)	39	42	1650	Brooks et al (5)
4 (1.8 - 3.1)	33 (27 - 47)	20 (9 – 29)	700 (269 - 1340)	Crosfill et al (8)
14 (Avg 5.3)	42 ± 7 (28.3 - 55.7)**	43.5 ± 8.1 (27.6 - 59.4)**	1791 ± 360 (1067 - 2514)**	Lees et al (15)
6 (Avg 2.7)	40 (31 – 52)	21.2 (8.8 - 29.1)	863 (311 - 1410)	Guyton (11)
6 (Avg 7.6)	1	1	1443 (1069 – 1855)	. Hayden (12)
21 (2.5 - 6.1)	26.4 ± 7.9 (24.1 - 28.6)**	27.7 ± 6.9 (26.7 - 28.7)**	709 ± 207 $(651 - 767)**$	Kelly et al (14)

All values are means + 1 standard deviation with ranges in parentheses except **.

^{*} No. of observations = 45.

^{** 95%} confidence limit of mean (1.96 x standard error of the mean).

TABLE 7--Comparison of values obtained by authors and other investigators for selected pulmonary RQ and θ_2 consumption in rhesus monkeys

7					
Condition of monkeys	No. of monkeys (wt. in kg)	FRC (m1)	Dead space (m1)	Dynamic pulmonary compliance $(m1/cm\ H_20)$	<pre>Expiratory flow (ml/sec)</pre>
Anesthetízed	9* (3.5 - 6.6)	112.7 ± 82 (73.6 - 205)	112.7 ± 82 5.68 ± 36 (73.6 - 205) (-16.2 - 11.2)	8.3 ± 4.3 (2.7 - 19.9)	61.8 ± 12.3 (23.4 - 115.9)
	11 (3.1 - 5.4)	1	1	1	
	4 (1.8 - 3.1)	87.5 (61 - 115)	1	12.3 (7.1 – 20.2)	
	14 (Avg 5.8)	1	12.6 ± 26.5 (-1.3 - 26.5)**	1	I
	11 · (2.7 - 3.6)	1	I	1	1
	8 (Avg 7.6)	1	1	1	69.1 ± 20.9 (53.6 - 84.6)**
	21 (2.5 - 6.1)	1	1	15.2 ± 10.0 $(7.8 - 22.6)**$	
Conscious	8		-	10.3 ± 2.9	;

All values are means + 1 standard deviation in parentheses except **. *No. of observation = 45, except dead space, which is 30. **95% confidence limit of mean (1.96 x standard error of the mean).

(2.4 - 4.3)

(5.2 - 14)

TABLE 7--Concluded

Dynamic pulmonary resistance (cm $H_2O/L/\sec$)	RQ	0 ₂ consumption (m1/min)	Transpulmonary pressure (cm H ₂ 0)	Reference
31.4 ± 23.4 (2.8 - 108.4)	0.78 ± 0.54 (0.45 - 1.38)	59.0 ± 37.2 (24.9 - 93)	5.3 ± 2.2 (2.1 - 11.9)	Present study
1	1	1	1	Brooks et al (5)
ſ	1	1	1	Crosfill et al (8)
1	0.77 ± 0.11	55.2 ± 49.3	;	Lees et al (15)
	(0.72 - 0.82)**	(28.6 - 81.8)**		
1	0.75 ± 0.02	22.7 ± 2.61	ŀ	Rakieten (24)
	(0.73 - 0.75)**	(21.1 - 24.3)**		
1	1	32.2	1	Hayden (12)
24.8 <u>+</u> 13.7 (14.6 - 34.9)**	l	1	1	Kelly et al (14)
22.0 ± 11.0 (10 - 44)	1	1	3.9 ± 0.6 (2.9 - 4.8)	Binns et al (3)

Figure Legends

- Fig 1--Three major parts of a head cover for a chaired conscious monkey.
- Fig 2--Equipment arrangement for studying pulmonary function in an anesthetized monkey.

References

- Binns, R., and Clark, G. C: An Experimental Model for the Assessment of the Effects of Cigarette Smoke Inhalation on Pulmonary Physiology. Ann Occup Hyg, 15, (1972):237-247.
- Binns, R., and Clark, G. C.: Lung Function in Cynomolgus Monkeys and Baboons. Folia Primatol, 17, (1972):209-217.
- 3. Binns, R., Clark, G. C., and Simpson, C. R.: Lung Function and Blood Gas Characteristics in the Rhesus Monkey. Lab Anim, 6, (1972): 189-198.
- 4. Bourne, G. H.: Collected Anatomical and Physiological Data from the Rhesus Monkey. In: <u>The Rhesus Monkey</u>, Vol. I. Edited by G. H. Bourne. Academic Fress, New York, NY (1975): 55-56.
- 5. Brooks, P. M., Richey, E. O., and Pickering, J. E.: Prompt Pulmonary Ventilation and Oxygen Consumption Changes in Rhesus Monkeys Associated with Whole-body Gamma-irradiation. Radiat Res, 6, (1957): 430-449.
- 6. Bruhn, J. M.: The Respiratory Metabolism of Infrahuman Primates.

 Am J Physiol, 110, (1934):477-484.
- 7. Chaffee, R. R. J., and Allen, J. R.: Studies on Metabolic Rate of Cold and Heat Acclimated Macaca mulatta. Physiologist, 13, (1970):163.
- 8. Crosfill, M. L., and Widdicombe, J. G.: Physical Characteristics of the Chest and Lungs and the Work of Breathing in Different Mammalian Species. J Physiol, 158, (1961):1-14.
- 9. Forsyth, R. P., Nies, A. S., Wyler, F., Neutze, J, and Melmon, K.
 L.: Normal Distribution of Cardiac Output in the Unanesthetized,
 Restrained Rhesus Monkey. J Appl Physiol, 25, (1968):736-741.

- 10. Giles, R. E., Finkel, M. P., and Mazurowski, J.: Use of an Analog On-Line Computer for the Evaluation of Pulmonary Resistance and Dynamic Compliance in the Anesthetized Dog. Arch Int. Pharmacodyn Ther, 194, (1971):213-222.
- 11. Guyton, A. C.: Measurement of the Respiratory Volumes of Laboratory Animals. Am J Physiol, 150, (1947):70-77.
- 12. Hayden, R. O.: Cardiopulmonary Measurements Obtained by Right and Left Cardiac Catheterization of <u>Macaca mulatta</u>. Technical Report AMRL-TDR-64-64, Office of Technical Services, US Department of Commerce, Washington, DC, (1964):1-6.
- 13. Huey, J. M., Martin, J., and Bryant, L. R.: Restraining Chair and Face Mask for Inhalation Toxicology Studies in Primates. J Med Primatol, 3, (1974):359-364.
- 14. Kelly, J. F., Cugell, D. W., Patterson, R., and Harris, K.E.:

 Acute Airway Obstruction in Rhesus Monkeys Induced by Pharmacologic

 and Immunologic Stimuli. J Lab Clin Med, 83, (1974):738-749.
- 15. Lees, M. H., Malinow, M. R., and Parer, J. T.: Cardiorespiratory

 Function of the Rhesus Monkey during Phencyclidine Analgesia. In

 Temas de Medicina. Edited by L. V. Decourt. Livro-Homenagem, Sao

 Paulo, Brazil, (1965):61-69.
- 16. Liu, C. T.: Cardiovascular and Renal Functions in Normal Rhesus Macaques. Am J Vet Res, 37, (1976):969-974.
- 17. Liu, C. T., DeLauter, R. D., and Griffin, M. J.: Pulmonary Dysfunction in Rhesus Monkeys during Staphylococcal B Enterotoxemia. Physiologist, 19, (1976):273.
- 18. Liu, C. T., and Higbee, G.: Determination of Body Surface Area in the Rhesus Monkey. J Appl Physiol 40, (1976):101-104.

- 19. Milic-Emili, J., Mead, J., Turner, J. M., and Glauser, E. M.:
 Improved Technique for Estimating Pleural Pressure from Esophageal
 Balloons. J Appl Physiol, 19, (1964):207-211.
- 20. Munson, E. S., Gillespie, J. R., and Wagman, I. H.:
 Respiratory Blood Gases and pH in Two Species of Unanesthetized
 Monkeys. J Appl Physiol, 28, (1970):108-109.
- 21. Patterson, R., and Booth, B. H.: Animal Models of Immediate-Type Respiratory Allergy. NY State J Med, 71, (1971):755-759.
- 22. Patterson, R., and Kelly, J. F.: Animal Models of the Asthmatic State. Ann Rev Med, 25, (1974):53-68.
- 23. Rakieten, N.: The Basal Heat Production of the Rhesus Monkey (Macaca mulatta). J Nutr, 10, (1935):357-362.

